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* * *	* *	* *	* *	* Welcome to STN International * * * * * * * * *		
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NEWS	5	FEB	02	Simultaneous left and right truncation (SLART) added for CERAB, COMPUAB, ELCOM, and SOLIDSTATE		
NEWS	6	FEB	02	GENBANK enhanced with SET PLURALS and SET SPELLING		
NEWS	7	FEB	0.6	Patent sequence location (PSL) data added to USGENE		
NEWS		FEB		COMPENDEX reloaded and enhanced		
NEWS	9	FEB	11	WTEXTILES reloaded and enhanced		
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NEWS	12	FEB	23	Several formats for image display and print options discontinued in USPATFULL and USPAT2		
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NEWS	14	FEB	23	TOXCENTER updates mirror those of MEDLINE - more precise author group fields and 2009 MeSH terms		
NEWS	15	FEB	23	Three million new patent records blast AEROSPACE into STN patent clusters		
NEWS	16	FEB	25	USGENE enhanced with patent family and legal status display data from INPADOCDB		
NEWS	17	MAR	06	INPADOCDB and INPAFAMDB enhanced with new display formats		
NEWS	18	MAR	11	EPFULL backfile enhanced with additional full-text applications and grants		
NEWS	19	MAR	11	ESBIOBASE reloaded and enhanced		
NEWS	20	MAR	20	CAS databases on STN enhanced with new super role for nanomaterial substances		
NEWS	21	MAR	23	CA/CAplus enhanced with more than 250,000 patent equivalents from China		
NEWS	2.2	MAR	3.0	IMSPATENTS reloaded and enhanced		
NEWS	23	APR	03	CAS coverage of exemplified prophetic substances enhanced		
NEWS	24	APR	07	STN is raising the limits on saved answers		

NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3, AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

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NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that specific topic.

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* * * * * * * * * * * * * * * * STN Columbus * * * * * * * * * * * * * * * * * *

FILE 'HOME' ENTERED AT 10:41:42 ON 21 APR 2009

=>

Uploading

THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE Do you want to switch to the Registry File?

Choice (Y/n):

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Some commands only work in certain files. For example, the EXPAND command can only be used to look at the index in a file which has an index. Enter "HELP COMMANDS" at an arrow prompt (=>) for a list of commands which can be used in this file.

=> FILE REGISTRY

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.22 0.22

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 10:41:57 ON 21 APR 2009
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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 19 APR 2009 HIGHEST RN 1136834-47-3

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TSCA INFORMATION NOW CURRENT THROUGH January 9, 2009.

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http://www.cas.org/support/stngen/stndoc/properties.html

=>

Uploading C:\Program Files\Stnexp\Queries\10587846.str

```
chain nodes:
6 7 9 11 12 15
ring nodes:
1 2 3 4 5
chain bonds:
1-15 2-11 5-6 6-7 6-9 11-12
ring bonds:
1-2 1-5 2-3 3-4 4-5
exact/norm bonds:
1-2 1-5 1-15 2-3 3-4 4-5 6-7 6-9 11-12
exact bonds:
2-2 1-5 6-6
isolated ring systems:
containing 1:
```

G1:Cb, Hy

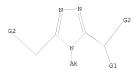
G2:Cy, Hy, Ph

Match level: 1:1Atom 3:Atom 4:Atom 5:Atom 6:CLASS 7:CLASS 9:CLASS 11:CLASS 12:CLASS 15:CLASS

L1 STRUCTURE UPLOADED

=> d 11

L1 HAS NO ANSWERS L1 STR



G1 Cb, Hy G2 Cv, Hy, Ph

Structure attributes must be viewed using STN Express query preparation.

=> s 11

SAMPLE SEARCH INITIATED 10:42:12 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 12855 TO ITERATE

15.6% PROCESSED 2000 ITERATIONS INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED) SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS: 250306 TO 263894
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s ll sss full

FULL SEARCH INITIATED 10:42:19 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 260501 TO ITERATE

83.5% PROCESSED 217493 ITERATIONS

0 ANSWERS 0 ANSWERS

0 ANSWERS

100.0% PROCESSED 260501 ITERATIONS

SEARCH TIME: 00.00.23

L3 0 SEA SSS FUL L1

--

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10587846.trn 04/21/2009

Page 4

```
chain nodes: 7 10 11 14 ring nodes: 1 2 3 4 5 6 15 16 chain bonds: 1 -14 2-10 5-6 6-7 10-11 ring bonds: 1 -14 2-10 5-6 6-7 10-11 ring bonds: 1 -2 1 -5 2-3 3-4 4-5 6-15 6-16 15-16 exact/norm bonds: 1 -2 1 -5 1-14 2-3 3-4 4-5 6-7 6-15 6-16 10-11 15-16 exact bonds: 2 -10 5-6 isolated ring systems: containing 1:
```

G1:Cb, Hy

G2:Cy, Hy, Ph

Match level: 1:Atom 2:Atom 4:Atom 5:Atom 6:CLASS 7:CLASS 10:CLASS 11:CLASS 14:CLASS 15:Atom 16:Atom

L4 STRUCTURE UPLOADED

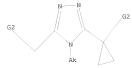
=> d 14

10587846.trn 04/21/2009

Page 5

L4 HAS NO ANSWERS

L4 STR



- G1 Cb, Hy
- G2 Cy, Hy, Ph

Structure attributes must be viewed using STN Express query preparation.

0 TO

0 ANSWERS

0 ANSWERS

=> s 14

SAMPLE SEARCH INITIATED 10:45:00 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 5094 TO ITERATE

39.3% PROCESSED 2000 ITERATIONS INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 97600 TO 106160

L5 0 SEA SSS SAM L4

PROJECTED ANSWERS:

=> s 14 sss full FULL SEARCH INITIATED 10:45:07 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 101216 TO ITERATE

100.0% PROCESSED 101216 ITERATIONS SEARCH TIME: 00.00.08

L6 0 SEA SSS FUL L4

=> Uploading C:\Program Files\Stnexp\Oueries\10587846b.str



```
9 10
ring nodes :
1 2 3 4 5 6 13 14 15 16 17 18 19 20
chain bonds :
2-9 5-6 6-16 9-10
ring bonds :
1-2 1-5 2-3 3-4 4-5 6-14 6-13 13-14 15-16 15-20 16-17 17-18 18-19
19-20
exact/norm bonds :
1-2 1-5 2-3 3-4 4-5 6-14 6-13 9-10 13-14
exact bonds :
2-9 5-6 6-16
normalized bonds :
15-16 15-20 16-17 17-18 18-19 19-20
isolated ring systems :
containing 1 :
```

G1:Cb,Hy

G2:Cy, Hy, Ph

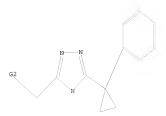
chain nodes :

Match level: 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:CLASS 9:CLASS 10:CLASS 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom

L7 STRUCTURE UPLOADED

=> d 17

L7 HAS NO ANSWERS L7 STR



G1 Cb,Hy G2 Cy,Hy,Ph

Structure attributes must be viewed using STN Express query preparation.

0 ANSWERS

0 ANSWERS

=> s 17

SAMPLE SEARCH INITIATED 10:46:52 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 73 TO ITERATE

100.0% PROCESSED 73 ITERATIONS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 948 TO 1972

PROJECTED ANSWERS:

WERS: 0 TO
0 SEA SSS SAM L7

=> s 17 sss full

FULL SEARCH INITIATED 10:46:59 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 1574 TO ITERATE

100.0% PROCESSED 1574 ITERATIONS

100.0% PROCESSED 1574 ITERATION SEARCH TIME: 00.00.01

L9 0 SEA SSS FUL L7

L8

Uploading C:\Program Files\Stnexp\Queries\10587846c.str

```
ring nodes : 1 2 3 4 5 6 10 11 12 13 14 15 16 17 18 19 20 21 22 23 chain bonds : 2-22 5-6 6-13 ring bonds : 1 2 1 3 14 15 16 17 18 19 20 21 22 23 chain bonds : 1 2 1-5 2-3 3-4 4-5 6-11 6-10 10-11 12-13 12-17 13-14 14-15 15-16 16-17 18-19 18-23 19-20 20-21 21-22 22-23 exact/norm bonds : 1 2 1-5 2-3 3-4 4-5 6-11 6-10 10-11 exact bonds : 2-22 5-6 6-13 normalized bonds : 1 2-12 12-12 12-17 13-14 14-15 15-16 16-17 18-19 18-23 19-20 20-21 21-22 22-23 isolated ring systems : containing 1 : 18 :
```

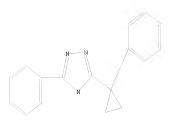
G1:Cb, Hy

G2:Cy, Hy, Ph

Match level: 1:Atcm 2:Atcm 3:Atcm 4:Atcm 5:Atcm 6:CLASS 10:Atcm 11:Atcm 12:Atcm 13:Atcm 14:Atcm 15:Atcm 16:Atcm 17:Atcm 18:Atcm 19:Atcm 20:Atcm 21:Atcm 22:Atcm 23:Atcm

L10 STRUCTURE UPLOADED

=> d 110 L10 HAS NO ANSWERS L10 STR



G1 Cb, Hy G2 Cy, Hy, Ph

Structure attributes must be viewed using STN Express query preparation.

=> s 110

SAMPLE SEARCH INITIATED 10:49:21 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -83 TO ITERATE

100.0% PROCESSED 83 ITERATIONS SEARCH TIME: 00.00.01

9 ANSWERS

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE** PROJECTED ITERATIONS: 1114 TO 2206 PROJECTED ANSWERS: 9 TO 360

L11 9 SEA SSS SAM L10

=> s 110 sss full

FULL SEARCH INITIATED 10:49:38 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 1699 TO ITERATE

100.0% PROCESSED 1699 ITERATIONS

209 ANSWERS

SEARCH TIME: 00.00.01

L12 209 SEA SSS FUL L10

Uploading C:\Program Files\Stnexp\Queries\10587846d.str

```
chain nodes :
24 25 26 27 28
ring nodes :
1 \quad \tilde{2} \quad 3 \quad 4 \quad 5 \quad 6 \quad 10 \quad 11 \quad 12 \quad 13 \quad 14 \quad 15 \quad 16 \quad 17 \quad 18 \quad 19 \quad 20 \quad 21 \quad 22 \quad 23
chain bonds :
2-22 5-6 6-13 19-24 24-25 25-26 25-28 26-27
ring bonds :
1-2 1-5 2-3 3-4 4-5 6-11 6-10 10-11 12-13 12-17 13-14 14-15 15-16
16-17 18-19 18-23 19-20 20-21 21-22 22-23
exact/norm bonds :
1-2 1-5 2-3 3-4 4-5 6-11 6-10 10-11 19-24 24-25 25-28 26-27
exact bonds :
2-22 5-6 6-13 25-26
normalized bonds :
12-13 12-17 13-14 14-15 15-16 16-17 18-19 18-23 19-20 20-21 21-22 22-23
isolated ring systems :
containing 1 : 18 :
```

G1:Cb, Hy

G2:Cy, Hy, Ph

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:CLASS 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:Atom 24:CLASS 25:CLASS 26:CLASS 27:CLASS 28:CLASS

L13 STRUCTURE UPLOADED

=> d 113

L13 HAS NO ANSWERS

L13 STR

Structure attributes must be viewed using STN Express query preparation.

=> s 113

G1 Cb, Hy G2 Cv, Hv, Ph

SAMPLE SEARCH INITIATED 10:52:07 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -13 TO ITERATE

100.0% PROCESSED 13 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE** **COMPLETE** BATCH PROJECTED ITERATIONS: 44 TO 476

0 SEA SSS SAM L13

PROJECTED ANSWERS: 0 TO

=> s 113 sss full

FULL SEARCH INITIATED 10:52:15 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 309 TO ITERATE

100.0% PROCESSED 309 ITERATIONS

5 ANSWERS SEARCH TIME: 00.00.01

L15 5 SEA SSS FUL L13

L14

Uploading C:\Program Files\Stnexp\Oueries\10587846e.str

```
24 26
ring nodes :
1 2 3 4 5 6 10 11 12 13 14 15 16 17 18 19 20 21 22 23
chain bonds :
1-26 2-22 5-6 6-13 19-24
ring bonds :
1-2 1-5 2-3 3-4 4-5 6-11 6-10 10-11 12-13 12-17 13-14 14-15 15-16
16-17 18-19 18-23 19-20 20-21 21-22 22-23
exact/norm bonds :
1-2 1-5 1-26 2-3 3-4 4-5 6-11 6-10 10-11 19-24
exact bonds :
2-22 5-6 6-13
normalized bonds :
12-13 12-17 13-14 14-15 15-16 16-17 18-19 18-23 19-20 20-21 21-22 22-23
isolated ring systems :
containing 1 : 18 :
```

G1:Cb, Hy

G2:Cv, Hv, Ph

chain nodes :

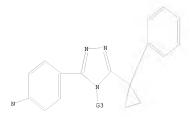
G3:Cb, Hy

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:Atom 24:CLASS 26:CLASS

L16 STRUCTURE UPLOADED

=> d 116 L16 HAS NO ANSWERS L16 STR



G1 Cb, Hy G2 Cy, Hy, Ph G3 Cb, Hy

Structure attributes must be viewed using STN Express query preparation.

=> s 116

SAMPLE SEARCH INITIATED 10:54:47 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 20 TO ITERATE

100.0% PROCESSED 20 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 132 TO 668
PROJECTED ANSWERS: 0 TO 0

L17 0 SEA SSS SAM L16

=> s 116 sss full

FULL SEARCH INITIATED 10:54:54 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 413 TO ITERATE

100.0% PROCESSED 413 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

L18 0 SEA SSS FUL L16

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```
27 ring nodes:
1 2 3 4 5 6 10 11 12 13 14 15 16 17 18 19 20 21 22 23 chain bonds:
1 2 5 2-22 5-6 6-13 ring bonds:
1-25 2-22 5-6 6-13 ring bonds:
1-2 1-5 2-3 3-4 4-5 6-11 6-10 10-11 12-13 12-17 13-14 14-15 15-16 16-17 18-19 18-23 19-20 20-21 21-22 22-23 exact/norm bonds:
1-2 1-5 1-25 2-3 3-4 4-5 6-11 6-10 10-11 exact bonds:
1-2 1-5 1-25 2-3 3-4 4-5 6-11 6-10 10-11 exact bonds:
2-22 5-6 6-13 normalized bonds:
12-13 12-17 13-14 14-15 15-16 16-17 18-19 18-23 19-20 20-21 21-22 22-23 isolated ring systems: containing 1: 18:
```

G1:Cb, Hy

G2:Cv, Hv, Ph

chain nodes :

G3:Cb, Hy

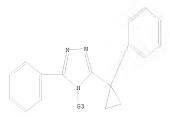
Match level: 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 14:Atom 15:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:Atom 25:CLASS

L19 STRUCTURE UPLOADED

=> d 119

L19 HAS NO ANSWERS

L19 STR



G1 Cb, Hv

G2 Cy, Hy, Ph

G3 Cb, Hy

Structure attributes must be viewed using STN Express query preparation.

=> s 119

SAMPLE SEARCH INITIATED 10:57:25 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 83 TO ITERATE

100.0% PROCESSED 83 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 1114 TO 2206

PROJECTED ITERATIONS: 1114 TO 220
PROJECTED ANSWERS: 0 TO

L20 0 SEA SSS SAM L19

=> s 119 sss full

FULL SEARCH INITIATED 10:57:31 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 1699 TO ITERATE

100.0% PROCESSED 1699 ITERATIONS SEARCH TIME: 00.00.01

L21 0 SEA SSS FUL L19

D21 0 35A 333 FOL 51

=> FIL HCAPLUS

 COST IN U.S. DOLLARS
 SINCE FILE
 TOTAL

 FULL ESTIMATED COST
 1310.76
 1310.76

0 ANSWERS

FILE 'HCAPLUS' ENTERED AT 10:58:19 ON 21 APR 2009
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FILE COVERS 1907 - 21 Apr 2009 VOL 150 ISS 17
FILE LAST UPDATED: 20 Apr 2009 (20090420/ED)
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HCAplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2008.

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d his

```
(FILE 'HOME' ENTERED AT 10:41:42 ON 21 APR 2009)
```

```
FILE 'REGISTRY' ENTERED AT 10:41:57 ON 21 APR 2009
L1
               STRUCTURE UPLOADED
L2
             0 S L1
L3
             0 S L1 SSS FULL
               STRUCTURE UPLOADED
L4
L5
             0 S L4
L6
             0 S L4 SSS FULL
L7
              STRUCTURE UPLOADED
L8
            0 S L7
L9
            0 S L7 SSS FULL
1.10
               STRUCTURE UPLOADED
L11
            9 S L10
L12
          209 S L10 SSS FULL
L13
               STRUCTURE UPLOADED
            0 S L13
L14
            5 S L13 SSS FULL
L15
L16
               STRUCTURE UPLOADED
            0 S L16
L17
L18
            0 S L16 SSS FULL
L19
              STRUCTURE UPLOADED
L20
            0 S L19
            0 S L19 SSS FULL
```

FILE 'HCAPLUS' ENTERED AT 10:58:19 ON 21 APR 2009

=> s 112

L22 5 L12

=> s 115 L23 1 L15

=> d 122 ibib abs hitstr tot

L22 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2008:1186333 HCAPLUS

DOCUMENT NUMBER: 149:548657

TITLE: Distinctive molecular inhibition mechanisms for selective inhibitors of human 11 β -hydroxysteroid

dehydrogenase type 1
AUTHOR(S): Tu, Hua; Powers, Jay P.; Liu, Jinsong; Ursu, Stefania;

Sudom, Athena; Yan, Xuelei; Xu, Haoda; Meininger, David; DeGraffenreid, Michael; He, Xiao; Jaen, Juan C.; Sun, Daqing; Labelle, Marc; Yamamoto, Hiroshi; Shan, Bei; Walker, Nigel P. C.; Wang, Zhulun

CORPORATE SOURCE: Department of Metabolic Disorders, Amgen Inc., South San Francisco, CA, 94080, USA

SOURCE: Bioorganic & Medicinal Chemistry (2008), 16(19), 8922-8931

CODEN: BMECEP; ISSN: 0968-0896

PUBLISHER: Elsevier Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English

$$_{\mathrm{F_{3}C}}$$
 $_{\mathrm{O}}$ $_{\mathrm{P_{r}-i}}^{\mathrm{N-N}}$ $_{\mathrm{II}}$

AB 11 β -Hydroxysteroid dehydrogenase type 1 (11 β -HSD1) catalyzes the NADPH dependent interconversion of inactive cortisone to active cortisol. Excess 11 β -HSD1 or cortisol leads to insulin resistance and metabolic syndrome in animal models and in humans. Inhibiting 11 β -HSD1

activity signifies a promising therapeutic strategy in the treatment of Type 2 diabetes and related diseases. Herein, the authors report two highly potent and selective small mol. inhibitors of human $11\beta-HSD1$. While compound (I), a sulfonamide, functions as a simple substrate competitive inhibitor, compound (II), a triazole, shows the kinetic profile of a mixed inhibitor. Co-crystal structures reveal that both compds. occupy the 11B-HSD1 catalytic site, but present distinct mol. interactions with the protein. Strikingly, compound (II) interacts much closer to the cofactor NADP+ and likely modifies its binding. Together, the structural and kinetic analyses demonstrate two distinctive mol. inhibition mechanisms, providing valuable information for future inhibitor design.

1080025-70-2P

RL: BSU (Biological study, unclassified); DMA (Drug mechanism of action); PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation and distinctive mol. inhibition mechanisms for selective inhibitors of human 11β-hydroxysteroid dehydrogenase type 1 and possible use for treatment of type 2 diabetes)

1080025-70-2 HCAPLUS RN

CN 4H-1,2,4-Triazole, 3-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-5-[4-(trifluoromethoxy)phenyl]- (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

29 L22 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2005:569372 HCAPLUS

DOCUMENT NUMBER: 143:97369

TITLE: Preparation of triazoles and related compounds as 11β-hydroxysteroid dehydrogenase 1 inhibitors

INVENTOR(S): Yamashita, Toshiro; Noda, Masakuni; Kawamoto,

Tomohiro; Irie, Kazuvuki

PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 65 pp. CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE JP 2005170939 20050630 JP 2004-337016 20041122 GI

PRIORITY APPLN. INFO .: OTHER SOURCE(S):

MARPAT 143:97369

JP 2003-391476 A 20031120

TT

Ι

Title compds. I [R1 = H, (un)substituted cyclic group; R2 = (un) substituted cyclic group; Ar = optionally furthermore substituted 5-, 6-membered heterocyclic ring; L1, L2 = bond, etc.] were prepared For example, benzylation of 4-ethyl-5-heptyl-2,4-dihydro-3H-1,2,4-triazol-3thione, e.g., prepared from octanoyl hydrazide in 2 steps, with 2,6-dichlorobenzyl chloride afforded compound II. In 11βHSD1 (11β-hydroxysteroid dehydrogenase type 1) inhibition assays, the IC50 value of compound II was 39 nM. Of note, compds. I are useful for the treatment of diabetes. Formulations are given. 856701-52-5P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(preparation of triazoles and related compds. as 11β-hydroxysteroid dehydrogenase 1 inhibitors)

RN 856701-52-5 HCAPLUS

4H-1,2,4-Triazole, 3-phenvl-5-(1-phenvlcvclopropvl)-4-(2-phenvlethvl)-(CA INDEX NAME)

CN

L22 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:423718 HCAPLUS

DOCUMENT NUMBER: 142:482046

TITLE: Preparation of triazole compounds as

11β-hydroxysteroid dehydrogenase 1 inhibitors INVENTOR(S): Cardozo, Mario G.; Powers, Jay P.; Goto, Hiroyuki; Harada, Kazuhito; Imamura, Katsuaki; Kakutani, Makoto;

Matsuda, Isamu; Ohe, Yasuhiro; Yata, Shinji Amgen SF LLC, USA; Japan Tobacco, Inc.

PATENT ASSIGNEE(S): PCT Int. Appl., 107 pp.

SOURCE: CODEN: PIXXD2

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| PATENT NO. | KIND DATE | APPLICATION NO. | DATE |
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| WO 2005044192
WO 2005044192 | | WO 2004-US35805 | 20041027 |
| W: AE, AG, AL, | AM, AT, AU, AZ, | BA, BB, BG, BR, BW, | BY, BZ, CA, CH, |
| CN, CO, CR, | CU, CZ, DE, DK, | DM, DZ, EC, EE, EG, | ES, FI, GB, GD, |
| GE, GH, GM, | HR, HU, ID, IL, | IN, IS, JP, KE, KG, | KP, KR, KZ, LC, |
| LK, LR, LS, | LT, LU, LV, MA, | MD, MG, MK, MN, MW, | MX, MZ, NA, NI, |
| NO, NZ, OM, | PG, PH, PL, PT, | RO, RU, SC, SD, SE, | SG, SK, SL, SY, |
| TJ, TM, TN, | TR, TT, TZ, UA, | UG, US, UZ, VC, VN, | YU, ZA, ZM, ZW |
| RW: BW, GH, GM, | KE, LS, MW, MZ, | NA, SD, SL, SZ, TZ, | UG, ZM, ZW, AM, |
| AZ, BY, KG, | KZ, MD, RU, TJ, | TM, AT, BE, BG, CH, | CY, CZ, DE, DK, |
| EE, ES, FI, | FR, GB, GR, HU, | IE, IT, LU, MC, NL, | PL, PT, RO, SE, |
| SI, SK, TR, | BF, BJ, CF, CG, | CI, CM, GA, GN, GQ, | GW, ML, MR, NE, |
| SN, TD, TG | | | |
| AU 2004286836 | A1 20050519 | AU 2004-286836 | 20041027 |
| CA 2543602 | A1 20050519 | CA 2004-2543602 | 20041027 |
| EP 1680114 | A2 20060719 | EP 2004-796647 | 20041027 |
| R: AT, BE, CH, | DE, DK, ES, FR, | GB, GR, IT, LI, LU, | NL, SE, MC, PT, |
| | | CZ, EE, HU, PL, SK | |
| | | JP 2006-538245 | |
| | | MX 2006-4674 | |
| | | US 2006-587846 | |
| PRIORITY APPLN. INFO.: | | US 2003-515537P | P 20031028 |
| | | WO 2004-US35805 | W 20041027 |
| OTHER SOURCE(S):
GI | CASREACT 142:48 | 2046; MARPAT 142:4820 |)46 |

AR The present invention provides triazole compds, of the following formula (I)or prodrugs thereof or pharmaceutically acceptable salts thereof [R1 = (un) substituted alkyl or cycloalkyl; Y = each (un) substituted cycloalkyl or heterocycloalkyl; Ar1 = aryl, heteroaryl; R2, R3 = H, halo, haloalkyl, alkyl group, (CH2)nOH, -N(R9)(R10), cyano, NO2, alkoxy, cycloalkyl, alkenyl, COR11, each (un)substituted arvl or heteroarvl group [wherein R9, R10 = H, alkvl,alkvlcarbonvl; R11 = OH, alkoxv, alkvl, (un)substituted NH2; n = 0-3]; Z = [CH(R14)]p, [CH(R14)]p-N(R16)[CH(R15)]q, each (un) substituted cycloalkylidene or heterocycloalkylidene (wherein p, q = 0-3; R14, R15 = group listed in R9 and R10]; Ar2 = aryl, heteroaryl, Q, Q1, Q2 [wherein X1 = (CH2)t; t = 0-2; V1 = :CH, :N; W1 = (un)substitutedCH2, O, S, SO2, SO, CO, (un) substituted NH]; R4, R5 = H, halo, OH, NO2, cyano, alkyl, alkoxy, COR27, SO2R27, each (un) substituted CONH2 or NH2; R27 = OH, alkoxy, alkyl, NH2, alkylamino, dialkylamino]. These triazole compds. are 11β-hydroxysteroid dehydrogenase 1-(11β-HSD1 or HSD1) and useful as therapeutic drugs for the treatment of diabetes, obesity or metabolic syndrome. Thus, Me N-methyl-4-phenylpiperidine-1-imidethiocarboxylate hydroiodide (452 mg) and 1-phenylcyclopropane carbohydrazide (176 mg) were suspended in 1,4-dioxane (2 mL) and water (0.4 mL) and sodium acetate (98 mg) were added and the mixture was heated under reflux overnight to give, after workup and silica gel chromatog., 117 mg 1-[4-methyl-5-(1-phenylcyclopropyl)-4H-[1,2,4]triazol-3-yl]-4phenylpiperidine hydrochloride (II). II showed IC50 of <10 nM against human HSD1. 1044957-17-6 1044957-18-7 1044957-19-8 1044957-20-1 1044957-21-2 1044957-22-3 1044957-23-4 1044957-24-5 1044957-25-6 1044957-46-1 1044957-47-2 1044957-48-3 1044957-49-4 1044957-50-7 1044957-51-8 1044957-55-2 1044957-56-3 1044957-65-4 1044957-67-6 1044957-68-7 1044957-69-8 1044957-70-1 1044957-71-2 1044957-72-3 1044957-73-4 1044957-74-5 1044957-75-6 1044957-76-7 1044957-77-8 1044957-78-9 1044957-79-0 1044957-80-3

(Preparation of triazole compounds as 11 β -hydroxysteroid dehyddrogenase 1 inhibitors) RN 1044957-17-6 HCAPLUS

CN INDEX NAME NOT YET ASSIGNED

RL: PRPH (Prophetic)

CN INDEX NAME NOT LET W221GNED

RN 1044957-18-7 HCAPLUS

CN INDEX NAME NOT YET ASSIGNED

- RN 1044957-19-8 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(3-fluorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)- (CA INDEX NAME)

- RN 1044957-20-1 HCAPLUS
- CN Benzenesulfonamide, 4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & F \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ &$$

- RN 1044957-21-2 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(2,4-dichlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)- (CA INDEX NAME)

RN 1044957-22-3 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

$$\begin{array}{c} \text{O} \\ \text{H}_2\text{N}-\text{C}-\text{CH}_2-\text{O} \\ \text{N} \\ \text{N}-\text{N} \end{array}$$

RN 1044957-23-4 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-24-5 HCAPLUS

CN 4H-1,2,4-Triazole, 3-(3-chlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)- (CA INDEX NAME)

RN 1044957-25-6 HCAPLUS

CN 4H-1, 2, 4-Triazole, 3-(3, 5-dichlorophenyl)-5-[1-(4fluorophenyl)cyclopropyl]-4-(1-methylethyl)- (CA INDEX NAME)

RN 1044957-46-1 HCAPLUS

CN INDEX NAME NOT YET ASSIGNED

RN

1044957-47-2 HCAPLUS Benzenamine, 4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-CN 1,2,4-triazol-3-v1]- (CA INDEX NAME)

RN 1044957-48-3 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-49-4 HCAPLUS

CN Acetamide, N-[2-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenoxy]ethyl]- (CA INDEX NAME)

RN 1044957-50-7 HCAPLUS

CN INDEX NAME NOT YET ASSIGNED

- RN 1044957-51-8 HCAPLUS
- CN INDEX NAME NOT YET ASSIGNED

- 1044957-55-2 HCAPLUS RN
- CN INDEX NAME NOT YET ASSIGNED

- HC1
- RN 1044957-56-3 HCAPLUS
- 4H-1,2,4-Triazole, 3-(2,3-dichlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl- (CA INDEX NAME) CN

- RN 1044957-65-4 HCAPLUS
- INDEX NAME NOT YET ASSIGNED CN

RN 1044957-67-6 HCAPLUS

CN INDEX NAME NOT YET ASSIGNED

RN 1044957-68-7 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-69-8 HCAPLUS

CN INDEX NAME NOT YET ASSIGNED

RN 1044957-70-1 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-71-2 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-72-3 HCAPLUS
CN Benzenamine, 3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H1,2,4-triazol-3-yl]-N-methyl-N-(1-methylethyl)- (CA INDEX NAME)

RN 1044957-73-4 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-74-5 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-75-6 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

- RN 1044957-76-7 HCAPLUS
- CN INDEX NAME NOT YET ASSIGNED

AcNH-CH2

- RN 1044957-77-8 HCAPLUS
- CN Benzenemethanamine, 3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

н2№-сн2

- RN 1044957-78-9 HCAPLUS
- CN Benzamide, 3-chloro-4-[5-(1-phenylcyclopropy1)-4-(2,2,2-trifluoroethy1)-4H-1,2,4-triazol-3-y1]- (CA INDEX NAME)

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RN 1044957-79-0 HCAPLUS CN INDEX NAME NOT YET ASSIGNED

RN 1044957-80-3 HCAPLUS CN 4H-1,2,4-Triazole, 3-(4-fluorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)- (CA INDEX NAME)

IT 851765-22-5P 851765-23-6P 851765-24-7P 851765-25-8P 851765-26-9P 851765-27-0P 851765-28-1P 851765-30-5P 851765-30-5P 851765-31-6P 851765-32-7P 851765-31-8P 851765-34-9P 851765-34-9P 851765-34-9P 851765-34-9P 851765-34-9P 851765-34-9P 851765-34-9P 851765-34-9P 851765-40-7P 851765-40-7P 851765-40-7P 851765-42-9P 851765-40-7P 851765-44-1P 851765-45-2P 851765-43-9P 851765-47-4P 851765-39-4P 851765-50-9P 851765-50-9P 851765-55-4P 851765-55-4P 851765-59-8P 851765-59-8P 851765-56-2-3P 851765-56-28-3P 851765-66-28-3P 851765-66-39 851765-66-

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851765-64-5P 851765-65-6P 851765-67-8P
851765-68-9P 851765-70-3P 851765-72-5P
851765-74-7P 851765-76-9P 851765-78-1P
851765-80-5P 851765-81-6P 851765-82-7P
851765-83-8P 851765-84-9P 851765-85-0P
851765-86-1P 851765-87-2P 851765-88-3P
851765-89-4P 851765-90-7P 851765-91-8P
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851766-38-6P 851766-40-0P 851766-41-1P
851766-42-2P 851766-43-3P 851766-44-4P
851766-45-5P 851766-46-6P 851766-47-7P
851766-48-8P 851766-49-9P 851766-50-2P
851766-52-4P 851766-53-5P 851766-54-6P
851766-55-7P 851766-56-8P 851766-57-9P
851766-58-0P 851766-59-1P 851766-60-4P
851766-61-5P 851766-62-6P 851766-63-7P
851766-64-8P 851766-65-9P 851766-66-0P
851767-63-0P 851767-64-1P 851767-66-3P
851767-67-4P 851767-68-5P 851768-01-9P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)
   (preparation of triazole compds. as 11\beta-hydroxysteroid dehydrogenase 1
   inhibitors for treatment of diabetes, obesity or metabolic syndrome)
851765-22-5 HCAPLUS
4H-1,2,4-Triazole, 3-(2,4-dichlorophenyl)-4-methyl-5-(1-phenylcyclopropyl)-
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RN

CN

HC1

RN 851765-23-6 HCAPLUS

CN 4H-1,2,4-Triazole, 3-(2,4-dichlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-, hydrochloride (1:1) (CA INDEX NAME)

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, hydrochloride (1:1) (CA INDEX NAME)

● HCl

- RN 851765-24-7 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(2-chloro-4-nitrophenyl)-4-methyl-5-(1-phenylcyclopropyl)- (CA INDEX NAME)

- RN 851765-25-8 HCAPLUS
- CN Acetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:1) (CA INDEX NAME)

HC1

- RN 851765-26-9 HCAPLUS
- CN Methanesulfonamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

- RN 851765-27-0 HCAPLUS
- CN Acetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]phenyl]-2-hydroxy- (CA INDEX NAME)

- RN 851765-28-1 HCAPLUS
- CN Acetamide, N-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-yl]phenyl]-2-methoxy- (CA INDEX NAME)

- RN 851765-29-2 HCAPLUS
- CN Acetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-2-(dimethylamino)-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

RN 851765-30-5 HCAPLUS
CN Benzenamine, 3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

- RN 851765-31-6 HCAPLUS
- CN 2-Pyrrolidinone, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

- RN 851765-32-7 HCAPLUS
- CN Butanamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

RN 851765-33-8 HCAPLUS

CN Propanamide, N-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-yl]phenyl]-2-methyl- (CA INDEX NAME)

RN 851765-34-9 HCAPLUS

CN 4-Morpholinecarboxamide, N-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

RN 851765-35-0 HCAPLUS

CN 4-Morpholineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851765-36-1 HCAPLUS
- CN 1-Piperidinecarboxylic acid, 4-[[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]amino]-, 1,1-dimethylethyl ester (CA INDEX NAME)

- RN 851765-37-2 HCAPLUS
- CN Urea, N'-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-N,N-dimethyl- (CA INDEX NAME)

- RN 851765-38-3 HCAPLUS
- CN Sulfamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:1) (CA INDEX NAME)

$$\begin{array}{c|c} & \text{Me} \\ & \text{N} \\ & \text{N} \\ & \text{N} \\ & \text{N} \end{array}$$

- RN 851765-39-4 HCAPLUS
- CN Urea, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3yl]phenyl]- (CA INDEX NAME)

- RN 851765-40-7 HCAPLUS
- CN 2-Oxazolidinone, 3-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

- RN 851765-41-8 HCAPLUS
- CN Ethanol, 2-[[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]amino]- (CA INDEX NAME)

HO- CH2- CH2- NH

- RN 851765-42-9 HCAPLUS
- CN Carbamic acid, [3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, ethyl ester (9CI) (CA INDEX NAME)

- RN 851765-43-0 HCAPLUS
- CN 2-Imidazolidinone, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

- RN 851765-44-1 HCAPLUS
- CN 1-Piperidinecarboxamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-methoxy- (CA INDEX NAME)

- RN 851765-45-2 HCAPLUS
- CN 1-Piperidinecarboxamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-3-hydroxy- (CA INDEX NAME)

RN 851765-46-3 HCAPLUS

CN 1-Piperidinecarboxamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-hydroxy- (CA INDEX NAME)

RN 851765-47-4 HCAPLUS

CN 1-Piperidineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-yl]phenyl]-4-hydroxy- (CA INDEX NAME)

RN 851765-49-6 HCAPLUS

CN 1-Piperidineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-3-hydroxy- (CA INDEX NAME)

- RN 851765-50-9 HCAPLUS
- CN 1-Piperidineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-methoxy- (CA INDEX NAME)

- RN 851765-51-0 HCAPLUS
- CN Morpholine, 4-[4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

- RN 851765-52-1 HCAPLUS
- CN Acetamide, N-[3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

10587846

RN 851765-54-3 HCAPLUS

CN Piperazine, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]phenyl]-, hydrochloride (1:3) (CA INDEX NAME)

●3 HC1

RN 851765-55-4 HCAPLUS

CN Ethanone, 1-[4-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-1-piperazinyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

RN 851765-57-6 HCAPLUS

CN Piperazine, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-

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triazol-3-y1]pheny1]-4-(methylsulfony1)- (CA INDEX NAME)

- RN 851765-59-8 HCAPLUS
- CN 1-Propanone, 1-14-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl|phenyl]-1-piperazinyl]-2-methyl-, hydrochloride (1:2) (CA INDEX NAME)

- 2 HC1
- RN 851765-61-2 HCAPLUS
- CN 1-Piperazinecarboxamide, 4-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-N,N-dimethyl-, hydrochloride (1:2) (CA INDEX NAME)

10587846

●2 HC1

851765-62-3 HCAPLUS
Piperidine, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-CN triazol-3-yl]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

RN 851765-64-5 HCAPLUS

CN 4-Piperidinol, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851765-65-6 HCAPLUS
- CN Morpholine, 4-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851765-67-8 HCAPLUS
- CN Piperidine, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-methyl-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851765-68-9 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-[2-chloro-4-(1-pyrrolidinyl)phenyl]-4-methyl-5-(1-phenylcyclopropyl)-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851765-70-3 HCAPLUS
- CN 4-Piperidinecarboxylic acid, 1-[3-chloro-4-[4-methyl-5-(1-phenyloyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, methyl ester, hydrochloride (1:2) (CA INDEX NAME)

10587846

●2 HC1

RN 851765-72-5 HCAPLUS

CN 4-Piperidinecarboxylic acid, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

RN 851765-74-7 HCAPLUS

CN 4-Piperidinecarboxamide, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-N-methyl- (CA INDEX NAME)

RN 851765-76-9 HCAPLUS

CN 1,4-Dioxa-8-azaspiro[4.5]decane, 8-[3-chloro-4-[4-methy1-5-(1-

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phenylcyclopropy1)-4H-1,2,4-triazo1-3-y1]pheny1]- (CA INDEX NAME)

- RN 851765-78-1 HCAPLUS
- CN 4-Piperidinecarboxylic acid, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yllphenyl]-, 2-methoxyethyl ester, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851765-80-5 HCAPLUS
- CN Carbamic acid, [1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-piperidinyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

- RN 851765-81-6 HCAPLUS
- CN 4-Piperidinamine, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:3) (CA INDEX NAME)

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●3 HC1

- RN 851765-82-7 HCAPLUS
- CN Acetamide, N-[1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-piperidinyl]- (CA INDEX NAME)

- RN 851765-83-8 HCAPLUS
- CN 3-Piperidinol, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851765-84-9 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-[2-chloro-4-[(4-methoxyphenyl)methoxy]phenyl]-4-

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methy1-5-(1-phenylcyclopropy1)- (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{N} \\ \text{Ph} \\ \text{N-N} \\ \end{array}$$

- RN 851765-85-0 HCAPLUS
- CN Phenol, 3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-, monohydrochloride (9CI) (CA INDEX NAME)

- HCl
- RN 851765-86-1 HCAPLUS
- CN Acetamide, 2-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenoxy]-, hydrochloride (1:1) (CA INDEX NAME)

$$\begin{array}{c|c} & \text{Me} \\ \\ \text{O} \\ \\ \text{H}_2\text{N}-\text{C}-\text{CH}_2-\text{O} \\ \end{array} \\ \text{N}-\text{N} \quad \text{Ph} \\ \end{array}$$

- HCl
- RN 851765-87-2 HCAPLUS
- CN Benzoic acid, 3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]-, methyl ester, hydrochloride (1:1) (CA INDEX NAME)

HC1

RN 851765-88-3 HCAPLUS CN 4H-1,2,4-Triazole, 3-[2-chloro-4-(methylsulfonyl)phenyl]-4-methyl-5-(1-phenylcyclopropyl)-, hydrochloride (1:1) (CA INDEX NAME)

$$\begin{array}{c|c} & \text{Me} \\ \hline & \text{N} \\ \\ \text{Me} \\ & \text{N} \\ \end{array}$$

● HC1

- RN 851765-89-4 HCAPLUS CN Benzoic acid, 3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 851765-90-7 HCAPLUS CN Benzamide, 3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazo1-3y1]- (CA INDEX NAME)

$$\begin{array}{c|c} C1 & \text{Me} \\ \hline \\ N & \\ N-N & Ph \\ \hline \\ O & \\ \end{array}$$

RN 851765-91-8 HCAPLUS

CN Methanone, [3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-yl]phenyl]-4-morpholinyl-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 851765-92-9 HCAPLUS
CN Benzamide, 3-chloro-N-methyl-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]-, hydrochloride (1:1) (CA INDEX NAME)

HCl

RN 851765-93-0 HCAPLUS
CN 4H-1,2,4-Triazole, 3-(2-chloro-4-methoxyphenyl)-4-methyl-5-(1-phenylcyclopropyl)- (CA INDEX NAME)

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RN 851765-94-1 HCAPLUS

CN Benzamide, 3-chloro-N, N-dimethyl-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 851765-95-2 HCAPLUS

CN Benzamide, 3-chloro-N-(2-hydroxyethyl)-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

RN 851765-96-3 HCAPLUS

CN Benzamide, 3-chloro-N-(1-methylethyl)-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

- RN 851765-97-4 HCAPLUS
- CN Methanone, [3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-1-piperidinyl-, hydrochloride (1:1) (CA INDEX NAME)

- HCl
- RN 851765-98-5 HCAPLUS
- CN Methanone, [3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl](4-hydroxy-1-piperidinyl)- (CA INDEX NAME)

- RN 851765-99-6 HCAPLUS
- CN Benzamide, N-(2-amino-2-oxoethyl)-3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

- RN 851766-00-2 HCAPLUS
- CN Benzamide, 3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-N-(2,2,2-trifluoroethyl)-, hydrochloride (1:1) (CA INDEX NAME)

- HCl
- RN 851766-01-3 HCAPLUS
- CN Benzamide, N-[2-(acetylamino)ethyl]-3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-, hydrochloride (1:1) (CA INDEX NAME)

- RN 851766-02-4 HCAPLUS
- CN Benzamide, 3-chloro-N-(2-methoxyethyl)-4-[4-methyl-5-(1-phenylcyclopropyl)- 4H-1,2,4-triazol-3-yl]-, hydrochloride (1:1) (CA INDEX NAME)

RN 851766-03-5 HCAPLUS

CN Ethanone, 1-[4-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]benzoyl]-1-piperazinyl]-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 851766-04-6 HCAPLUS

CN Benzamide, 3-chloro-N-[2-(dimethylamino)ethyl]-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

RN 851766-05-7 HCAPLUS

CN Benzamide, 3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-y1]-N-[2-(4-morpholiny1)ethy1]- (CA INDEX NAME)

- RN 851766-06-8 HCAPLUS
- CN Benzenamine, 3-chloro-N,N-dimethyl-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-, hydrochloride (1:2) (CA INDEX NAME)

- ●2 HC1
- RN 851766-07-9 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(4-chloro-2-methoxyphenyl)-4-methyl-5-(1-phenylcyclopropyl)-, hydrochloride (1:1) (CA INDEX NAME)

- HCl
- RN 851766-08-0 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(2-chloro-4-fluorophenyl)-4-methyl-5-(1-phenylcyclopropyl)-, hydrochloride (1:1) (CA INDEX NAME)

RN 851766-09-1 HCAPLUS CN 4H-1,2,4-Triazole, 3-(4-chloro-2-nitrophenyl)-4-methyl-5-(1-phenylcyclopropyl)-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

- RN 851766-10-4 HCAPLUS
- CN Benzenamine, 5-chloro-2-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851766-11-5 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(4-chloro-2-fluorophenyl)-4-methyl-5-(1-phenylcyclopropyl)-, hydrochloride (1:1) (CA INDEX NAME)

HC1

- RN 851766-16-0 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(4-bromo-2-chlorophenyl)-4-methyl-5-(1phenylcyclopropyl)- (CA INDEX NAME)

- RN 851766-17-1 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(2-chlorophenyl)-4-methyl-5-(1-phenylcyclopropyl)-(CA INDEX NAME)

- RN 851766-18-2 HCAPLUS
- CN Benzenemethanol, 3-chloro-a-methyl-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-a-(trifluoromethyl)- (CA INDEX NAME)

- RN 851766-22-8 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(4-chlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-, hydrochloride (1:1) (CA INDEX NAME)

- HCl
- RN 851766-23-9 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(2,4-dichlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl)-4-methyl- (CA INDEX NAME)

- RN 851766-24-0 HCAPLUS
- CN 4-Piperidinemethanol, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851766-25-1 HCAPLUS
- CN 4-Piperidinecarboxamide, 1-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-y1]pheny1]- (CA INDEX NAME)

- RN 851766-26-2 HCAPLUS CN 4-Piperidinecarboxamide, 1-
 - CN 4-Piperidinecarboxamide, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-N-(2-methoxyethyl)- (CA INDEX NAME)

- RN 851766-29-5 HCAPLUS
- CN Methanone, [1-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-

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triazol-3-yl]phenyl]-4-piperidinyl]-1-piperidinyl- (CA INDEX NAME)

RN 851766-30-8 HCAPLUS
CN Methanone [1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1, 2, 4triazol-3-yl]phenyl]-4-piperidinyl]-4-morpholinyl- (CA INDEX NAME)

RN 851766-31-9 HCAPLUS CN 4-Piperidinecarboxamide, 1-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-N,N-dimethyl- (CA INDEX NAME)

RN 851766-32-0 HCAPLUS
CN 4-Piperidinecarboxamide, 1-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)4H-1,2,4-triazo1-3-y1]pheny1]-N-(2,2,2-trifluoroethy1)- (CA INDEX NAME)

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- RN 851766-33-1 HCAPLUS
- CN Urea, N-[1-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-yl]phenyl]-4-piperidinyl]-N'-methoxy- (CA INDEX NAME)

- RN 851766-34-2 HCAPLUS
- CN 2-Oxazolidinone, 3-[3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

- RN 851766-35-3 HCAPLUS
- CN Ethanol, 2-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenoxy]-, hydrochloride (1:1) (CA INDEX NAME)

RN 851766-36-4 HCAPLUS

CN Ethanol, 2-[[3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]phenyl]amino]- (CA INDEX NAME)

HO- CH2- CH2- NH

RN 851766-38-6 HCAPLUS

CN 4H-1,2,4-Triazole, 3-(2,5-dichlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 851766-40-0 HCAPLUS

CN Benzoic acid, 3-methoxy-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4triazol-3-yl]- (CA INDEX NAME)

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- RN 851766-41-1 HCAPLUS
- CN Benzamide, 3-methoxy-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

- RN 851766-42-2 HCAPLUS
- CN Benzoic acid, 3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]-, hydrochloride (1:1) (CA INDEX NAME)

$$\begin{picture}(200,0) \put(0,0){\line(1,0){100}} \put(0,0){\line(1,0){10$$

- HCl
- RN 851766-43-3 HCAPLUS
- CN Benzamide, 3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

- RN 851766-44-4 HCAPLUS
- CN Benzamide, 3-chloro-4-[5-[1-(4-fluoropheny1)cyclopropy1]-4-methy1-4H-1,2,4-triazol-3-y1]-N-methy1- (CA INDEX NAME)

- RN 851766-45-5 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-[2-chloro-4-(2,2,2-trifluoroethoxy)phenyl]-4-methyl-5-(1-phenylcyclopropyl)-, hydrochloride (1:1) (CA INDEX NAME)

Me

● HCl

- RN 851766-46-6 HCAPLUS
- CN Ethanamine, 2-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenoxy]-, hydrochloride (1:2) (CA INDEX NAME)

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●2 HC1

- RN 851766-47-7 HCAPLUS
- CN 3-Pyrrolidinol, 1-[3-chloro-4-[4-methy1-5-(1-phenylcyclopropy1)-4H-1,2,4-triazol-3-y1]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

●2 HC1

- RN 851766-48-8 HCAPLUS
- CN Benzamide, 4-[4-(1-methylethyl)-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3yl]- (CA INDEX NAME)

- RN 851766-49-9 HCAPLUS
- CN 4H-1,2,4-Triazole, 3-(2-chlorophenyl)-5-[1-(4-fluorophenyl)cyclopropyl]-4methyl- (CA INDEX NAME)

- RN 851766-50-2 HCAPLUS
- CN Urea, N'-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3yl]phenyl]-N-hydroxy-N-methyl- (CA INDEX NAME)

- RN 851766-52-4 HCAPLUS
- CN Benzamide, 4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-1,2,4triazol-3-yl]- (CA INDEX NAME)

- RN 851766-53-5 HCAPLUS
- CN 2-Oxazolidinone, 3-[4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

- RN 851766-54-6 HCAPLUS
- CN Carbamic acid, [4-chloro-3-[5-[1-(4-fluoropheny1)cyclopropy1]-4-methyl-4H-1,2,4-triazol-3-yl]phenyl]-, methyl ester (9CI) (CA INDEX NAME)

- RN 851766-55-7 HCAPLUS
- CN 2-Oxazolidinone, 3-[4-chloro-3-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

- RN 851766-56-8 HCAPLUS
- CN Urea, N-[3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-yl]phenyl]-N'-methoxy- (CA INDEX NAME)

RN 851766-57-9 HCAPLUS

CN Urea, N-[4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-1,2,4-triazol-3-yl]phenyl]-N'-methoxy- (CA INDEX NAME)

RN 851766-58-0 HCAPLUS

CN 2-Oxazolidinone, 3-[3-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-1,2,4-triazol-3-yl]phenyl]- (CA INDEX NAME)

RN 851766-59-1 HCAPLUS

CN Urea, N-[3-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-1,2,4triazol-3-yl]phenyl]-N'-methoxy- (CA INDEX NAME)

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- RN 851766-60-4 HCAPLUS
- CN Benzamide, 4-chloro-3-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4triazol-3-yl]- (CA INDEX NAME)

- RN 851766-61-5 HCAPLUS
- CN Benzamide, 4-chloro-3-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

- RN 851766-62-6 HCAPLUS
- CN Benzamide, 3-chloro-4-[4-ethyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

$$\begin{array}{c|c} C1 & Et \\ N & N \end{array}$$

- RN 851766-63-7 HCAPLUS
 CN Benzamide, 3-chloro-4-[4-ethyl-5-[1-(4-fluorophenyl)cyclopropyl]-4H-1,2,4triazol-3-yl]- (CA INDEX NAME)
 - C1 Et N N N
- RN 851766-64-8 HCAPLUS
- CN Benzamide, 3-[4-(1-methylethyl)-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

- RN 851766-65-9 HCAPLUS
- CN Benzamide, 3-[5-[1-(4-fluorophenyl)cyclopropyl]-4-(1-methylethyl)-4H-1,2,4triazol-3-yl]- (CA INDEX NAME)

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$$\begin{array}{c|c} & & & & F \\ & & & & \\ & & & \\ H_2N-C & & & N \\ & & & N-N \end{array}$$

- RN 851766-66-0 HCAPLUS
- CN Benzenemethanol, 4-chloro-α-methyl-3-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]-α-(trifluoromethyl)- (CA INDEX NAME)

- RN 851767-63-0 HCAPLUS
- CN Methanone, [3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-morpholinyl- (CA INDEX NAME)

- RN 851767-64-1 HCAPLUS
- CN Benzamide, 3-chloro-N-methyl-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

RN 851767-66-3 HCAPLUS

CN Benzamide, 3-chloro-N, N-dimethyl-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]- (CA INDEX NAME)

RN 851767-67-4 HCAPLUS

CN Methanone, [3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-1-piperidinyl- (CA INDEX NAME)

RN 851767-68-5 HCAPLUS

CN Ethanone, 1-[4-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]benzoyl]-1-piperazinyl]- (CA INDEX NAME)

851768-01-9 HCAPLUS RN

CN Morpholine, 4-[3-chloro-4-[5-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-4H-1,2,4-triazol-3-y1]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

2 HC1

REFERENCE COUNT: THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN

DOCUMENT NUMBER: 140:27832

2003:991491 HCAPLUS TITLE: Preparation of triazolvl 11B-hydroxysteroid dehydrogenase-1 inhibitors for the treatment of

diabetes, obesity and dyslipidemia

INVENTOR(S): Olson, Steven H.; Balkovec, James M.; Zhu, Yuping

PATENT ASSIGNEE(S): Merck & Co., Inc., USA PCT Int. Appl., 144 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

ACCESSION NUMBER:

PATENT NO. KIND DATE APPLICATION NO. DATE

| WO 2003104208 | | | | | A1 20031218 | | | | 2003 | -US17 | 20030606 | | | | | | | |
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| | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC | , EE | , ES, | FI, | GB, | GD | , GE, | GH, | |
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| | R: | | | | | | | | | | | | | | | , MC, | | |
| | | IE, | | | | | | | | | | | | | | , SK | | |
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| PRIORIT: | Y APP | LN. | INFO | . : | | | | | | | | | | | | 20020 | | |
| | | | | | | | | | | | | | | | | 20030 | | |
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| | | | | | | | | | | US | 2003 | -4576 | 82 | | A3 | 20030 | 609 | |
| OTHER SO | DURCE | (S): | | | MAR | PAT | 140: | 2783: | 2 | | | | | | | | | |

AB Title compds. I [A = halo, alkyl, Ph, etc.; B = H, halo, alkyl, S-alkyl, etc. or A, B = taken together are (un)substituted alkylene; R1 = H, OH, halo, alkyl, alkoxy, aryl, etc.; R2 = alkyl, alkoxy, Ph, etc.; R3 = alkyl, alkenyl, thioalkoxy, aryl, heterocyclyl, etc. or R2-3 = taken together fused 5-6-membered alkyl/aryl ringl are prepared For instance, 2,2-diphenylbutanoic acid is converted to the corresponding hydraxide (DMF, E13N, TFFH, H2NNH2, 0°, 30 min). 8-Methoxy-2,3,4,5,6,7-hexahydroazocine is then reacted with the intermediate (DMF, 120°, overnight) to give II. Example compds. exhibit IC50 < 500 nM for 11β -hydroxysteroid dehydrogenase-1 (11 β -HSDI). I are useful for the treatment of diabetes, such as noninsulin-dependent diabetes (NIDDM), hyperglycemia, obesity, insulin resistance, dylsipidemia, hyperlipidemia, hypertension, Syndrome X and other symptoms associated with NIDDM.

ΙI

IT 633317-12-1P 633317-13-2P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU

(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of triazolyl 11β-hydroxysteroid dehydrogenase-1 inhibitors for treatment of diabetes, obesity and dyslipidemia)

633317-12-1 HCAPLUS RN

4H-1,2,4-Triazole, 3-[1-(4-chlorophenyl)cyclopropyl]-4-methyl-5-[4-(trifluoromethoxy)phenvll- (CA INDEX NAME)

633317-13-2 HCAPLUS RN

CN 4H-1,2,4-Triazole, 3-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-5-[4-(trifluoromethoxy)phenyl]- (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

2 L22 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2003:991490 HCAPLUS

DOCUMENT NUMBER: 140:27831

TITLE: Preparation of triazolyl 11β-hydroxysteroid dehydrogenase-1 inhibitors for the treatment of

diabetes, obesity and dyslipidemia

Olson, Steven H.; Balkovec, James M.; Zhu, Yuping INVENTOR(S): PATENT ASSIGNEE(S): Merck & Co., Inc., USA

SOURCE: PCT Int. Appl., 91 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE WO 2003104207 A2 20031218 WO 2003-US17898 20030606

| WO | 2003 | 10420 | 07 | | A3 | | 2004 | 0325 | | | | | | | | | | |
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| | W: | AE, | AG, | AL, | AM, | AT, | AU, | AZ, | BA, | BB | , BG, | BR, | BY, | BZ, | CA | , CH, | CN, | |
| | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC | , EE, | ES, | ΓI, | GB, | GD | , GE, | GH, | |
| | | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KE | , KG, | KR, | KZ, | LC, | LK | , LR, | LS, | |
| | | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW | , MX, | MZ, | NI, | NO, | NZ | , OM, | PH, | |
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| | | UA, | UG, | US, | UZ, | VC, | VN, | YU, | ZA, | ZM | , ZW | | | | | | | |
| | RW: | GH, | GM, | KE, | LS, | MW, | MZ, | SD, | SL, | SZ | , TZ, | UG, | ZM, | ZW, | AM | , AZ, | BY, | |
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| | CN 1659151 | | | A 20050824 | | | | | CN | 20030606 | | | | | | | | |
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20030 | | |
| OTHER SO | DURCE | (S): | | | MARI | PAT | 140: | 2783 | | US | 2003- | 4010 | 02 | | мэ | 20030 | 005 | |

OTHER SOURCE(S):

MARPAI 140:2/031

AB Title compds. I [A = halo, alkyl, Ph, etc.; B = H, halo, alkyl, S-alkyl, etc. or A, B = taken together are (un)substituted alkylene; RI = H, OH, halo, alkyl, alkoxy, aryl, etc.; R2 = alkyl, alkoxy, Ph, etc.; R3 = alkyl, alkenyl, thioalkoxy, aryl, heterocyclyl, etc. or R2-3 = taken together fused 5-6-membered alkyl/aryl ringl are prepared For instance, 2,2-diphenylbutanoic acid is converted to the corresponding hydrazide (DMF, E13N, TFFH, H2NNH2, 0°, 30 min).
8-Methoxy-2,3,4,5,6,7-hexahydroazocine is then reacted with the intermediate (DMF, 120°, overnight) to give II. Example compds. exhibit IC50 < 500 nM for 11P-hydroxysteroid dehydrogenase-1 (11P-HSD1). I are useful for the treatment of diabetes, such as noninsulin-dependent diabetes (NIDM), hyperglycemia, obesity, insulin

Page 79

resistance, dylsipidemia, hyperlipidemia, hypertension, Syndrome X and other symptoms associated with NIDDM.

633317-12-1P 633317-13-2P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(preparation of triazolvl 11B-hydroxysteroid dehydrogenase-1 inhibitors for treatment of diabetes, obesity and dyslipidemia)

RN 633317-12-1 HCAPLUS

CN 4H-1,2,4-Triazole, 3-[1-(4-chlorophenyl)cyclopropyl]-4-methyl-5-[4-(trifluoromethoxy)phenyl] - (CA INDEX NAME)

633317-13-2 HCAPLUS

4H-1,2,4-Triazole, 3-[1-(4-fluorophenyl)cyclopropyl]-4-methyl-5-[4-(trifluoromethoxy)phenyl]- (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L23 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2009 ACS on STN

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ACCESSION NUMBER: 2005:423718 HCAPLUS

DOCUMENT NUMBER: 142:482046

TITLE: Preparation of triazole compounds as

11β-hvdroxysteroid dehvdrogenase 1 inhibitors Cardozo, Mario G.; Powers, Jay P.; Goto, Hiroyuki; INVENTOR(S):

Harada, Kazuhito; Imamura, Katsuaki; Kakutani, Makoto;

Matsuda, Isamu; Ohe, Yasuhiro; Yata, Shinji

PATENT ASSIGNEE(S): Amgen SF LLC, USA; Japan Tobacco, Inc.

SOURCE: PCT Int. Appl., 107 pp.

CODEN: PIXXD2

10587846.trn 04/21/2009

Page 80

DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND DATE | APPLICATION NO. | DATE |
|---|---|--|---|
| | | | |
| WO 2005044192
WO 2005044192 | | WO 2004-US35805 | 20041027 |
| CN, CO, CR,
GE, GH, GM,
LK, LR, LS,
NO, NZ, OM,
TJ, TM, TN,
RW: BW, GH, GM,
AZ, BY, KG, | CU, CZ, DE, DK,
HR, HU, ID, IL,
LT, LU, LV, MA,
PG, PH, PL, PT,
TR, TT, TZ, UA,
KE, LS, MW, MZ,
KZ, MD, RU, TJ, | BA, BB, BG, BR, BW, E
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MD, MG, MK, MN, MW, M
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NA, SD, SL, SZ, TZ, U
TM, AT, BE, BG, CH, C | S, FI, GB, GD, CP, KR, KZ, LC, KX, MZ, NA, NI, GG, SK, SL, SY, UU, ZA, ZM, ZW GG, ZM, ZW, AM, CY, CZ, DE, DK, |
| | BF, BJ, CF, CG, | IE, IT, LU, MC, NL, F
CI, CM, GA, GN, GQ, G | |
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| | | CA 2004-2543602 | |
| | | EP 2004-796647 | |
| | | GB, GR, IT, LI, LU, N | |
| | | CZ, EE, HU, PL, SK | ib, 55, iic, 11, |
| | | JP 2006-538245 | 20041027 |
| MV 2006004674 | 7 200/0413 | MX 2006-4674 | 20041027 |
| TIG 2000004074 | A 20001120 | US 2006-587846 | 20060420 |
| PRIORITY APPLN. INFO.: | | US 2003-515537P | |
| INTONITI ALPEN. INFO.: | | WO 2004-US35805 | |
| OTHER SOURCE(S): | CASREACT 142:482 | | |

The present invention provides triazole compds. of the following formula (I)or prodrugs thereof or pharmaceutically acceptable salts thereof [R1 = (un) substituted alkyl or cycloalkyl; Y = each (un) substituted cycloalkyl or heterocycloalkyl; Arl = aryl, heteroaryl; R2, R3 = H, halo, haloalkyl,

alkyl group, (CH2)nOH, -N(R9)(R10), cyano, NO2, alkoxy, cycloalkyl, alkenyl, COR11, each (un)substituted arvl or heteroarvl group [wherein R9, R10 = H, alkyl, alkylcarbonyl; R11 = OH, alkoxy, alkyl, (un) substituted NH2; n = 0-3]; Z = [CH(R14)]p, [CH(R14)]p-N(R16)[CH(R15)]q, each (un) substituted cycloalkylidene or heterocycloalkylidene (wherein p, q = 0-3; R14, R15 = group listed in R9 and R10]; Ar2 = aryl, heteroaryl, Q, 01, 02 [wherein X1 = (CH2)t; t = 0-2; V1 = :CH, :N; W1 = (un)substituted CH2, O, S, SO2, SO, CO, (un) substituted NH1; R4, R5 = H, halo, OH, NO2, cyano, alkyl, alkoxy, COR27, SO2R27, each (un) substituted CONH2 or NH2; R27 = OH, alkoxy, alkyl, NH2, alkylamino, dialkylamino]. These triazole compds. are 11β-hydroxysteroid dehydrogenase 1-(11β-HSD1 or HSD1) and useful as therapeutic drugs for the treatment of diabetes, obesity or metabolic syndrome. Thus, Me N-methyl-4-phenylpiperidine-1-imidethiocarboxylate hydroiodide (452 mg) and 1-phenylcyclopropane carbohydrazide (176 mg) were suspended in 1,4-dioxane (2 mL) and water (0.4 mL) and sodium acetate (98 mg) were added and the mixture was heated under reflux overnight to give, after workup and silica gel chromatog., 117 mg 1-[4-methvl-5-(1-phenvlcvclopropvl)-4H-[1,2,4]triazol-3-vl]-4phenylpiperidine hydrochloride (II). II showed IC50 of <10 nM against human HSD1.

IT 851765-35-0P 851765-47-4P 851765-49-6P 851765-50-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of triazole compds. as 11β-hydroxysteroid dehydrogenase 1 inhibitors for treatment of diabetes, obesity or metabolic syndrome)
RN 851765-35-0 HCAPLUS

CN 4-Morpholineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-, hydrochloride (1:2) (CA INDEX NAME)

● 2 HCl

RN 851765-47-4 HCAPLUS

CN 1-Piperidineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-hydroxy- (CA INDEX NAME)

RN 851765-49-6 HCAPLUS

CN 1-Piperidineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-3-hydroxy- (CA INDEX NAME)

RN 851765-50-9 HCAPLUS

CN 1-Piperidineacetamide, N-[3-chloro-4-[4-methyl-5-(1-phenylcyclopropyl)-4H-1,2,4-triazol-3-yl]phenyl]-4-methoxy- (CA INDEX NAME)

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REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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